Catalysis and covalent binding energies of antibodies

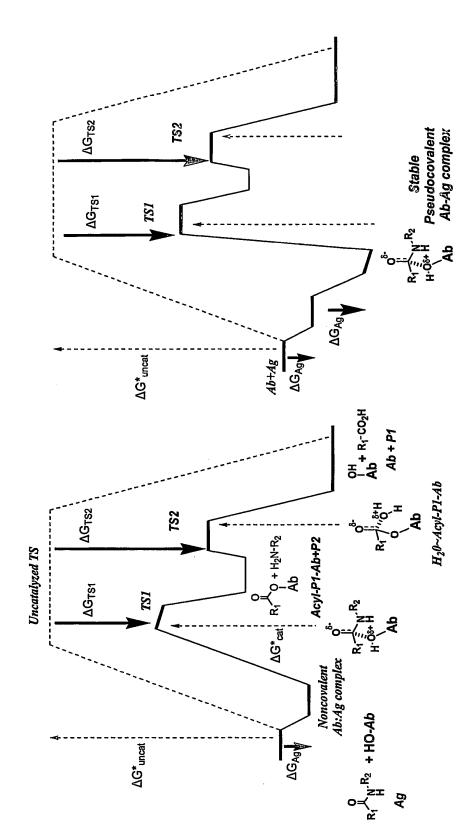
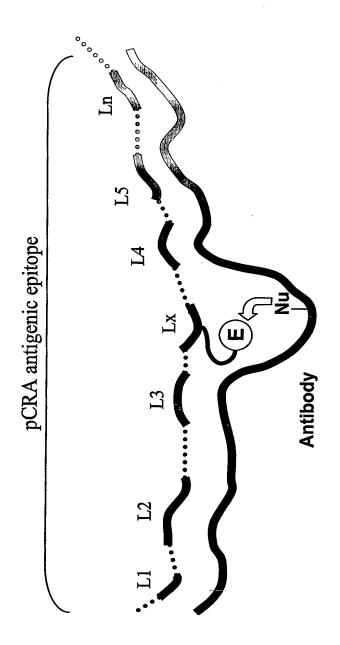


Fig 1

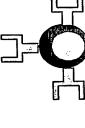
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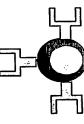


Induction of catalyst synthesis

Ordinary polypeptide antigen, Slow transmembrane signal

Aborted clonal selection, tolerance



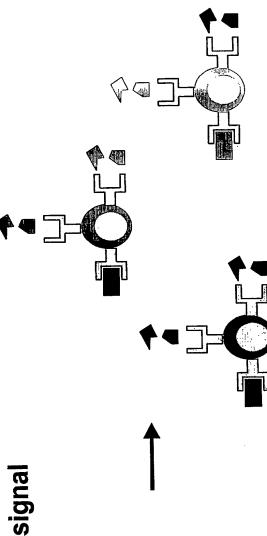






Increased turnover, proliferation

Rapid fransmembrane signal C⊚valent stimulation,



L' =sulfhydryl group of Cys

Fig 4

General structure of pCRA

• L'-Y"-Y'-Y, Example ∜

L' =
$$\beta$$
-carboxyl group of Asp
$$\begin{bmatrix} O = C \\ - C \\ NH \\ Y' = ethyl amine group \\ HC - CH_3 \\ HO - P - O \end{bmatrix}$$

• L'-Y"-Y'-Y, Example 3

• L'.Y".Y'.Y, Example 2

Fig 5A

(attached to Y")-H

Fig 5B

Fig $5\mathbb{C}$

Fig 5D

A. Electron withdrawing substituents with peptide extension

B. Electron donating substituents with peptide extension

Example of 6: DAIMP

Example of 5: 4'-Carboxybenzo-18-crown-6

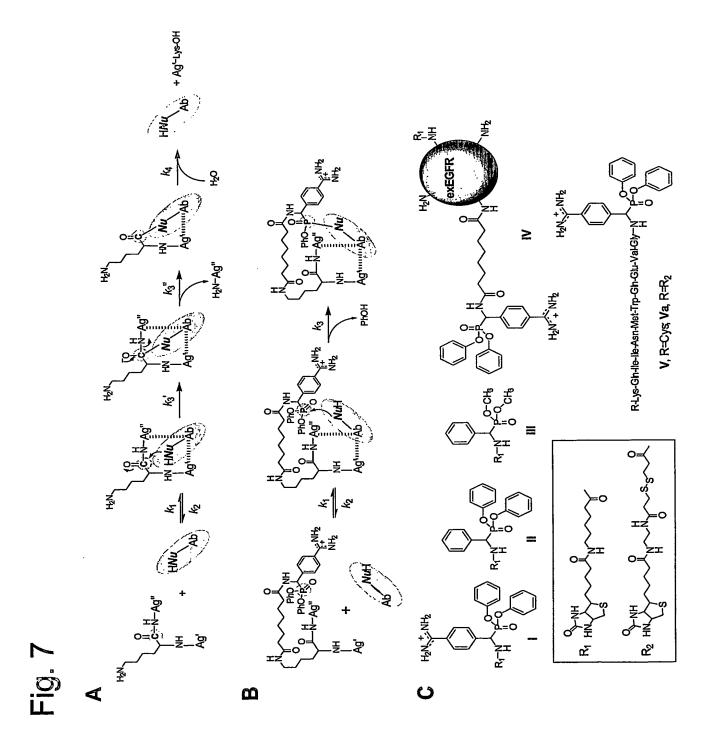
Example of 4: EDTA

Selected examples of metal binding moiety

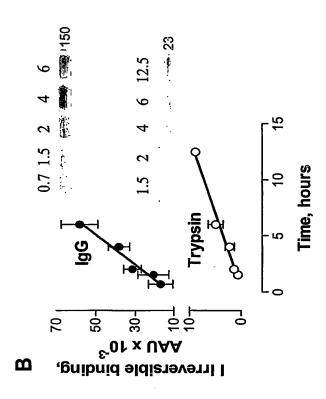
- domain-derived peptide
 3. (Cys-Aaa-Cys): metallothionein β-domain-derived peptide
 4. EDTA 2. (Cys-Aaa-Cys-Cys): metallothionein α-
- 5. a crown ether
- 6. DAMP

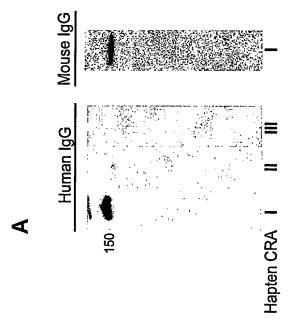
Example of 3: Cys-Ser-Cys H₂N-CHC-N-CHC-N-CHC-OH CH₂ CH₂ CH₂ CH₂ CH₂ CH₂ CH₂ CH₃ CH₂ CH₂ CH₃ CH₂ CH₂ CH₃ CH₃ CH₂ CH₃ CH₃ CH₃ Example of 2: Cys-Ser-Cys-Example of 1: (His),

Fig 6

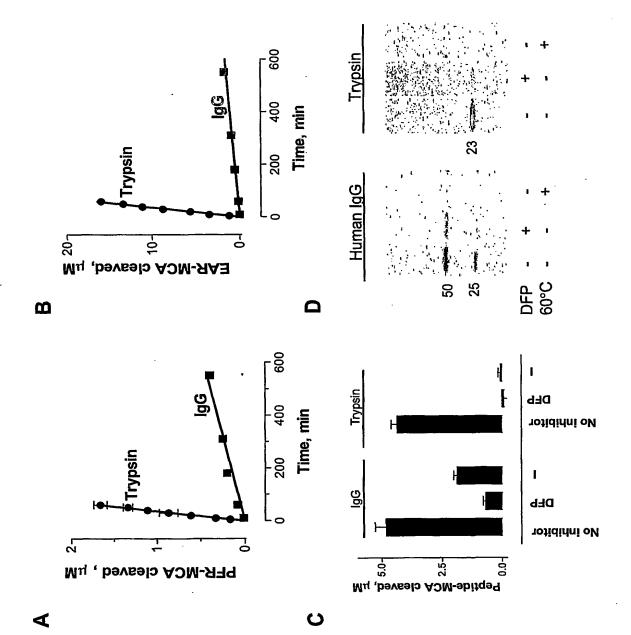




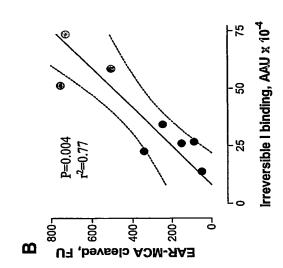




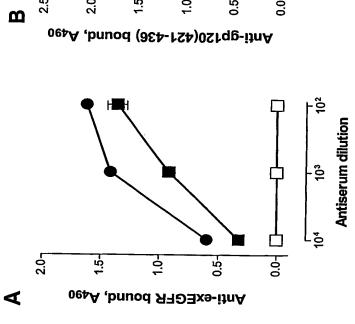


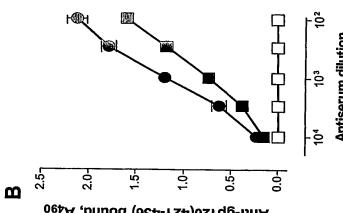


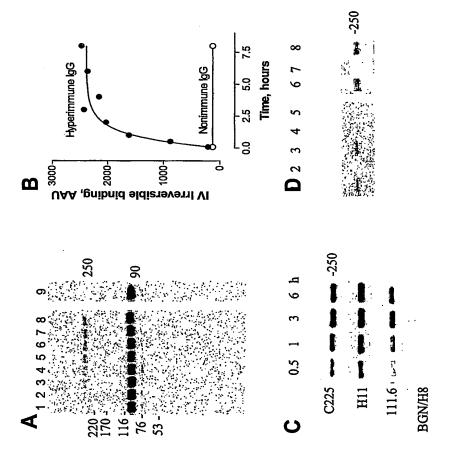












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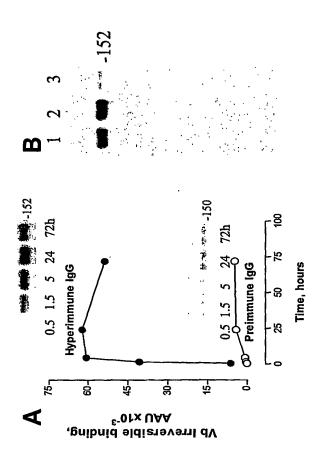
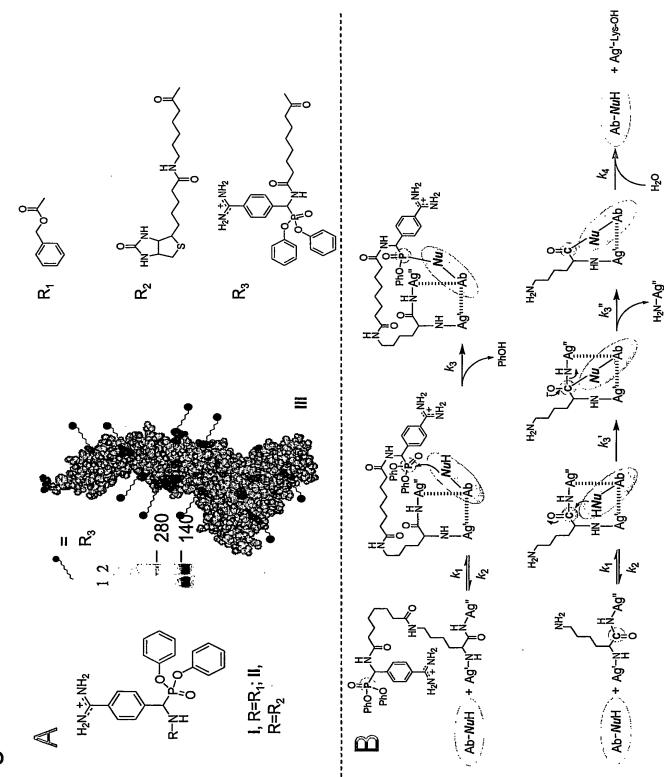
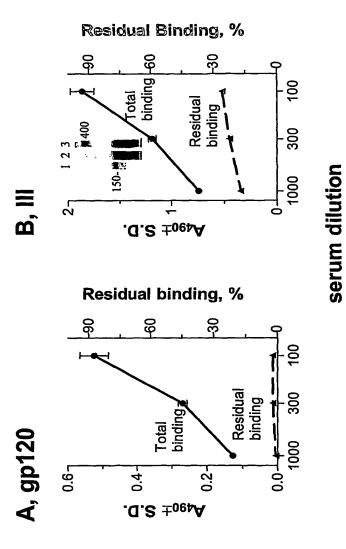


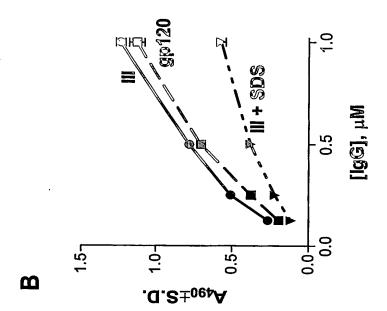
Fig 14

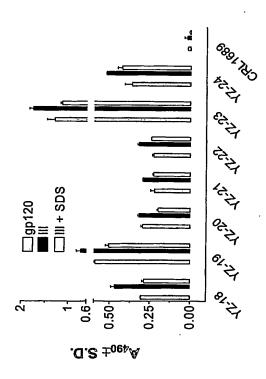




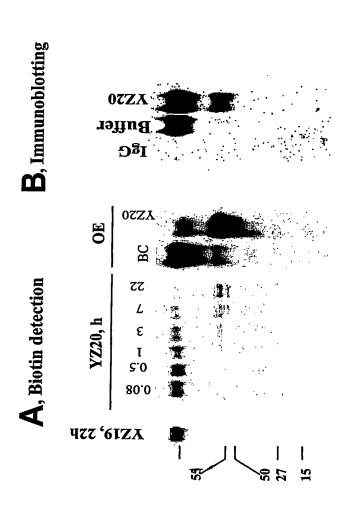


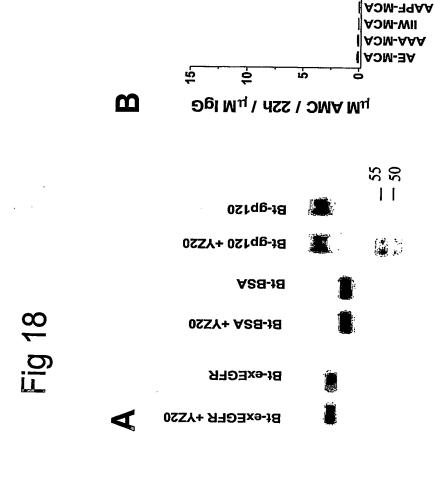


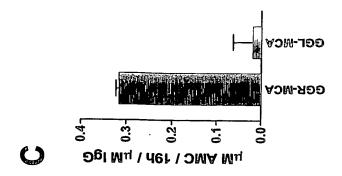












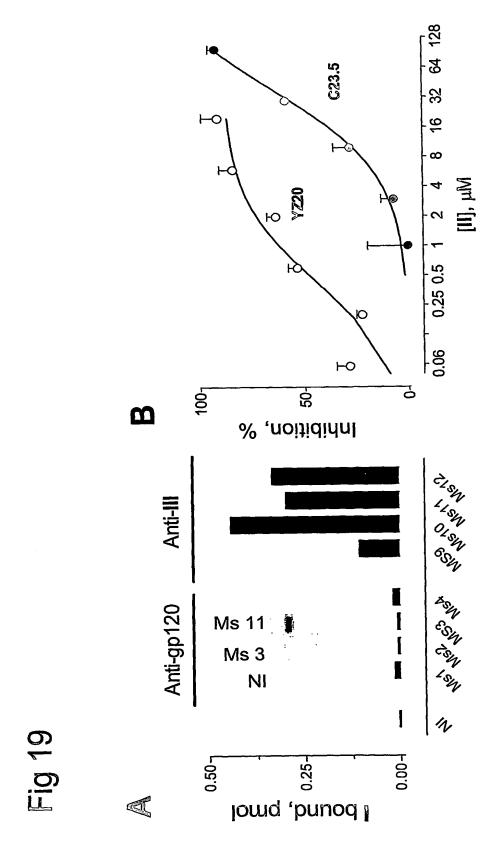
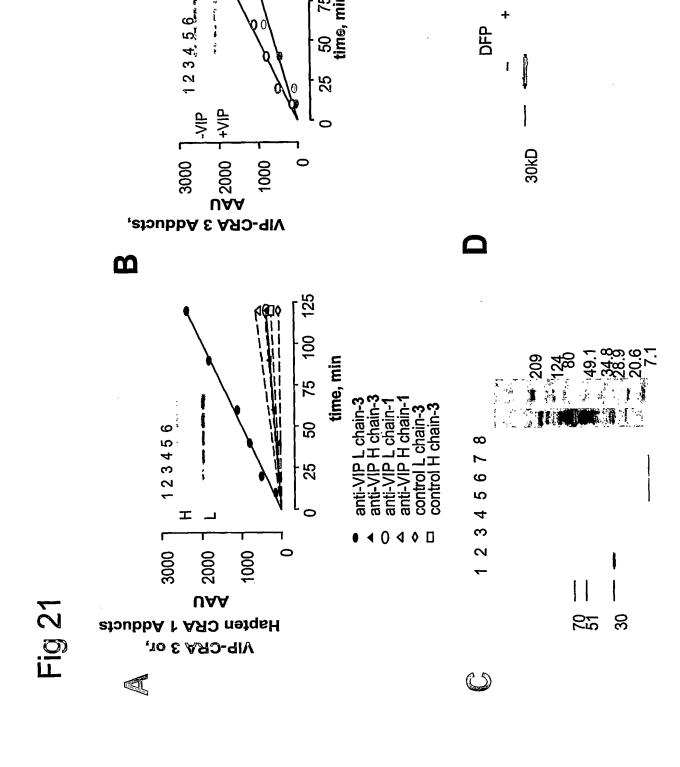


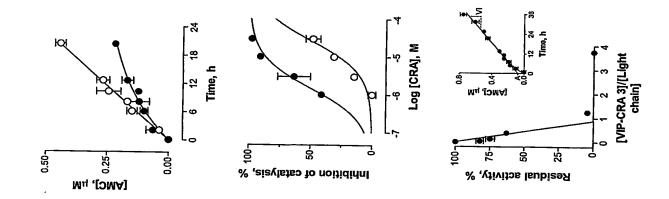
Fig 20

-Lys-Tyr-Leu-Asn-Ser-Ile-Leu-Asn-NH2 R-His-Ser-Asp-Ala-Val-Phe-Thr-Asp-Asn-Tyr-Thr-Arg-Leu-Arg-Lys-Gln-Met-Ala-Val--N H 3: R = D-biotinyl 1: R = 6-(D-biotinamido)hexanoyl 2: R = O-succinimidylsuberyl

Solid support R₁-His-Ser-Asp-Ala-Val-Phe-Thr-Asp-Ash-Tyr-Thr-Arg-Leu-Arg-Lys-Gin-Met-Ala-Val-Lys-Lys-Tyr-Leu-Asn-Ser-Ile-Leu-As Boc tBuO tBuO tBu Pmc Boc Tr 4b (R₁=D-biotinyl; R₂=Mtt) 4c (R₁=D-biotinyl; R₂=H) - 4a (R₁=Fmoc; R₂=Mtt) Fmoc-Asn(Trt)-Nv,v **≡**, .≥

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Fig 22

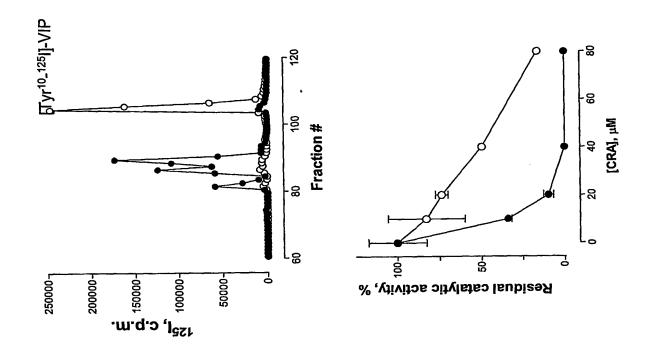
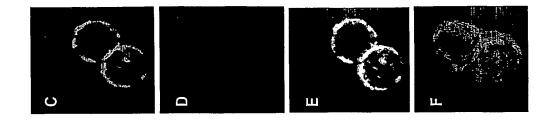


Fig 23

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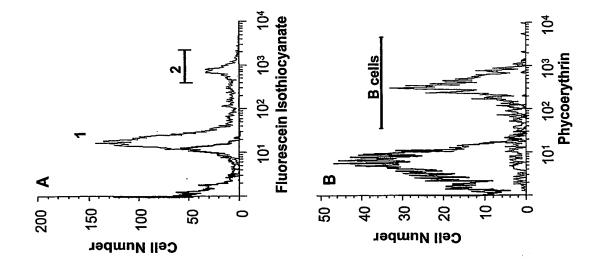


Fig 25

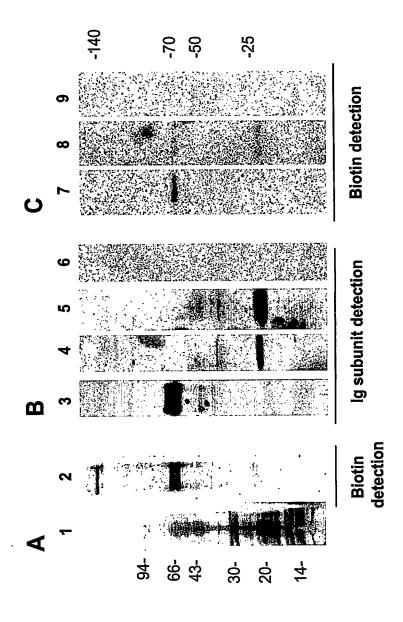
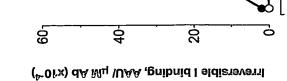
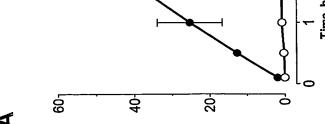
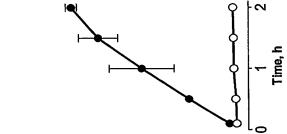


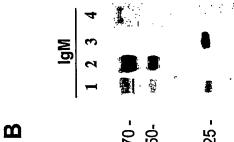
Fig 26



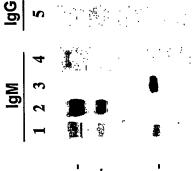


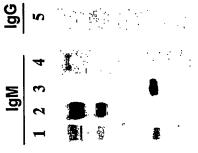






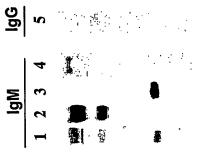
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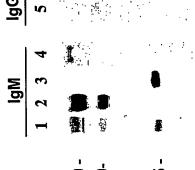


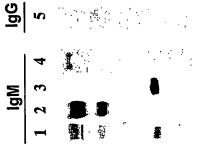


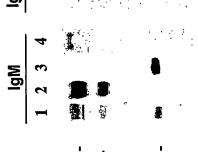
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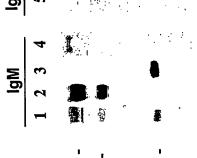
Velocity, AAU/h/µM Ab subunit (x10

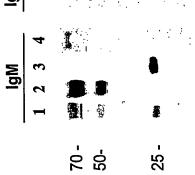














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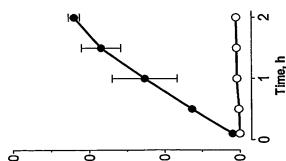
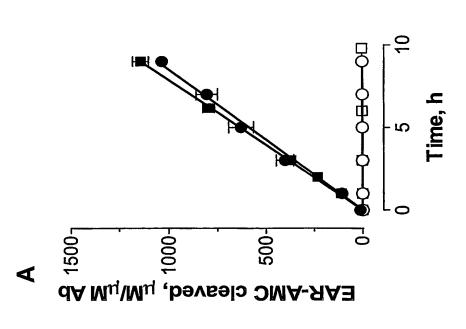
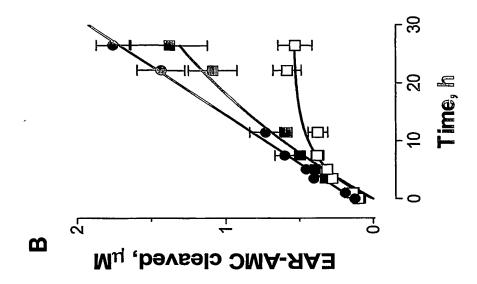
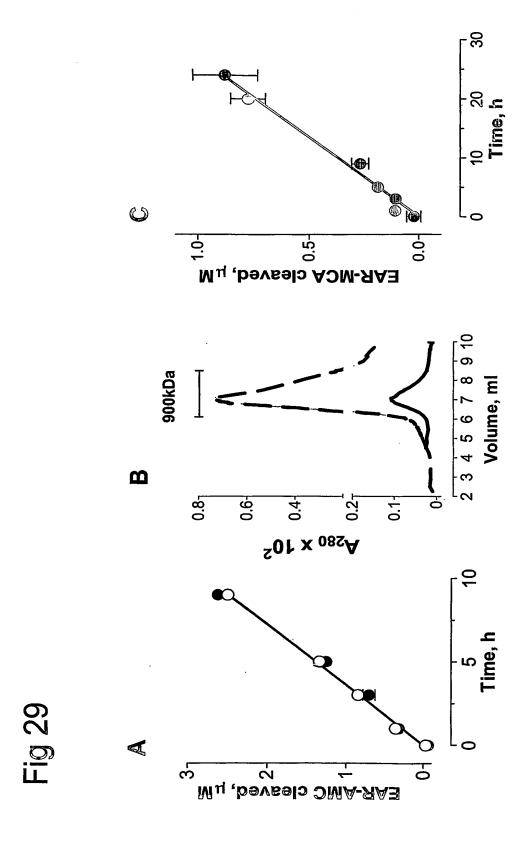


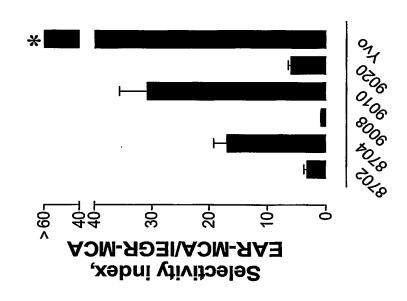
Fig 28



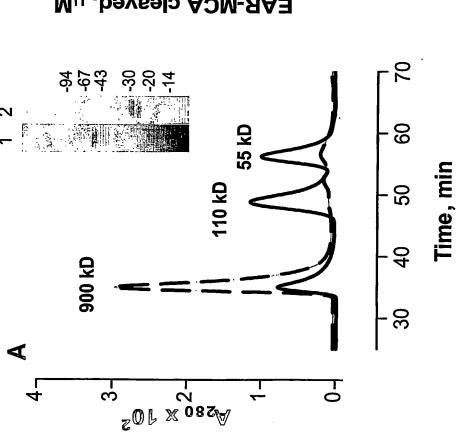


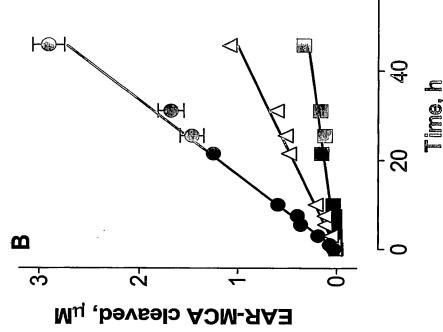


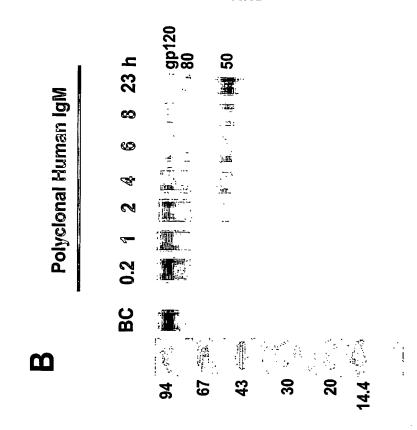


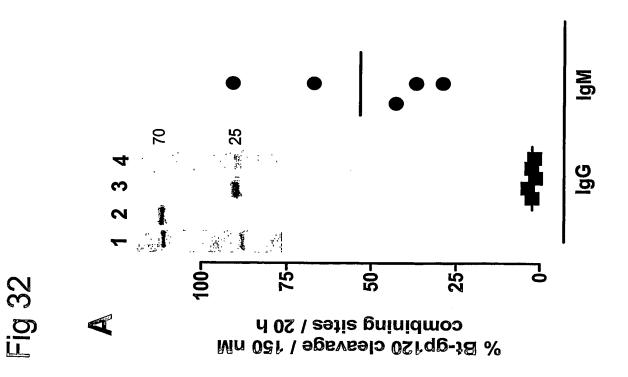












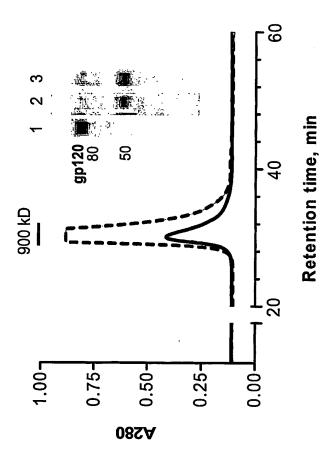
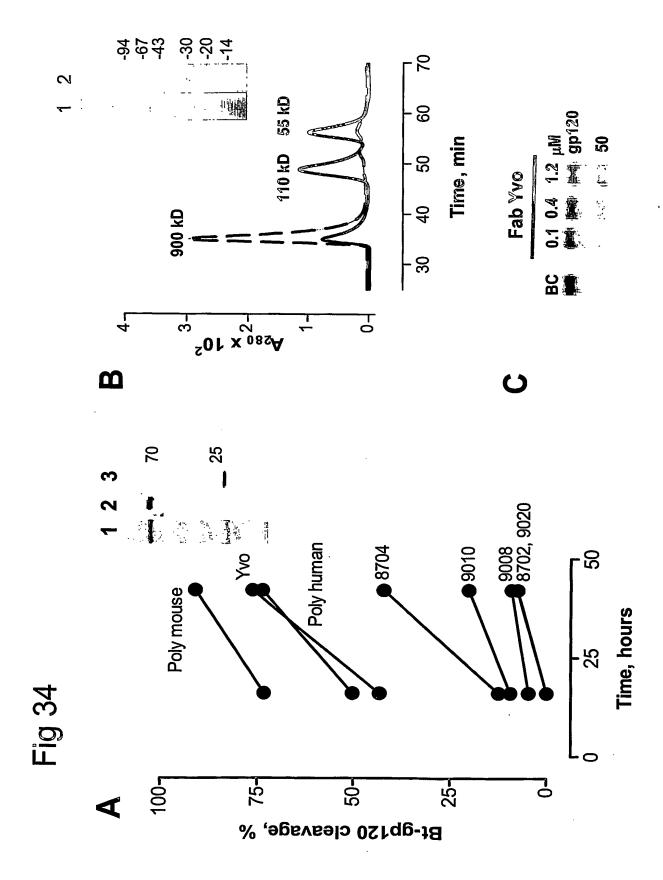
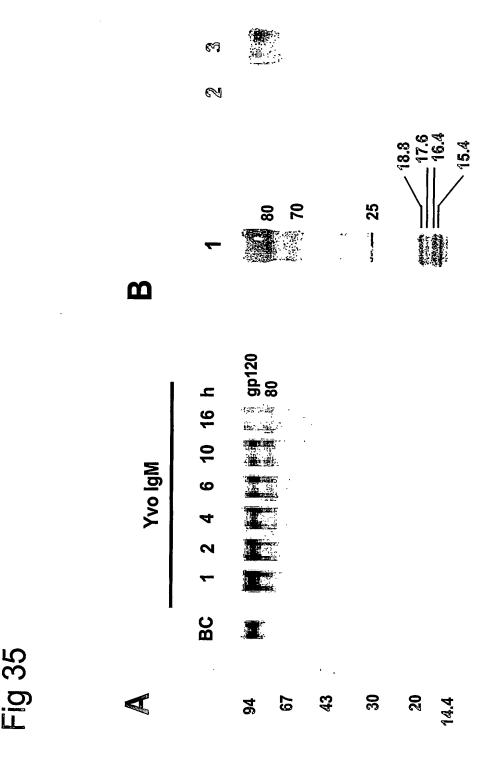
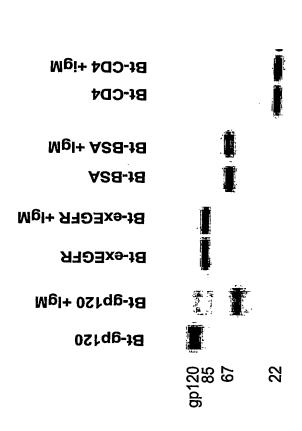


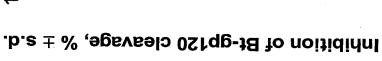
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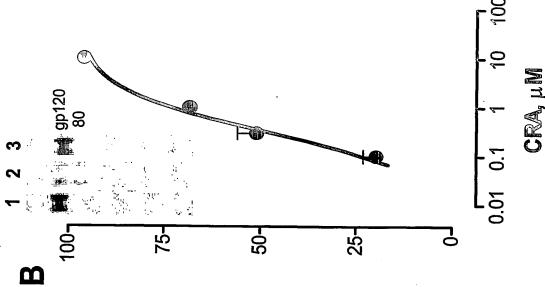


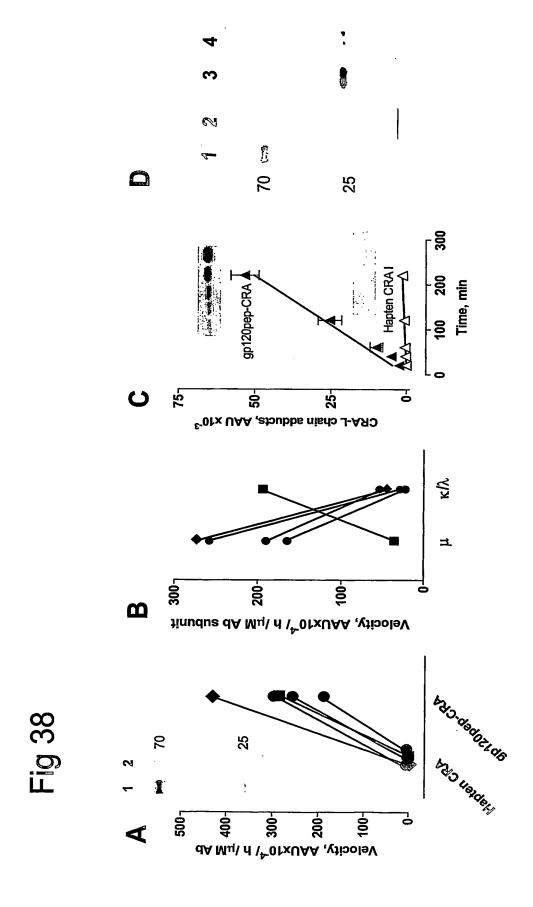


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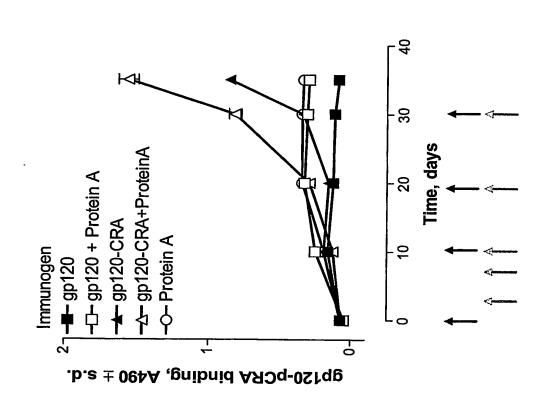


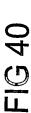


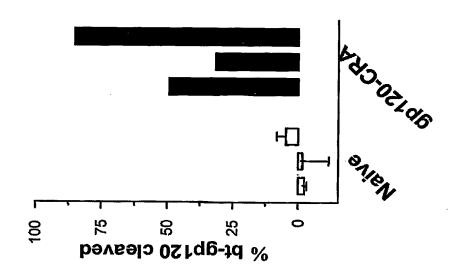




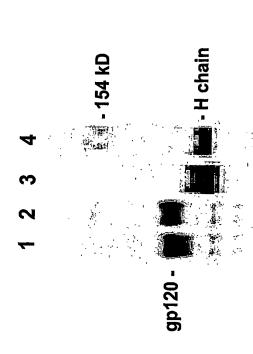












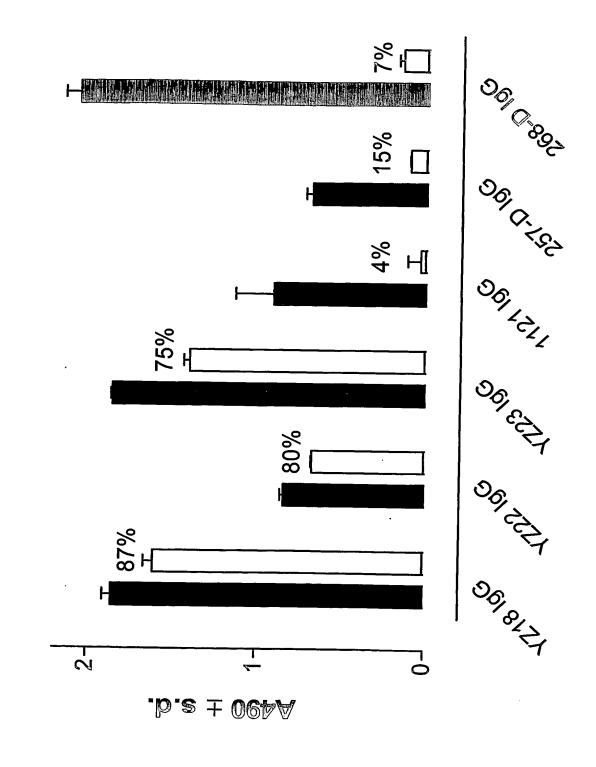
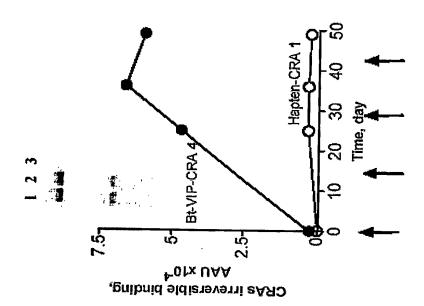
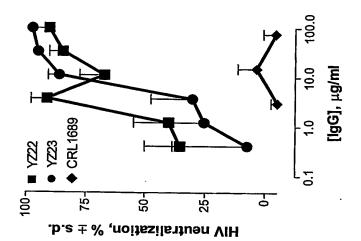


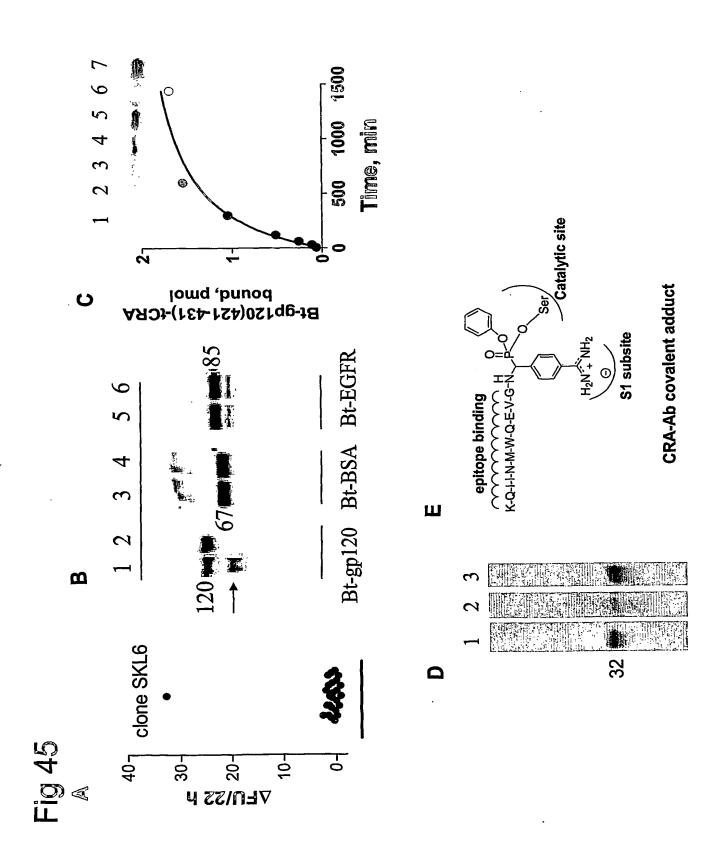
Fig 42

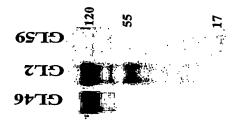




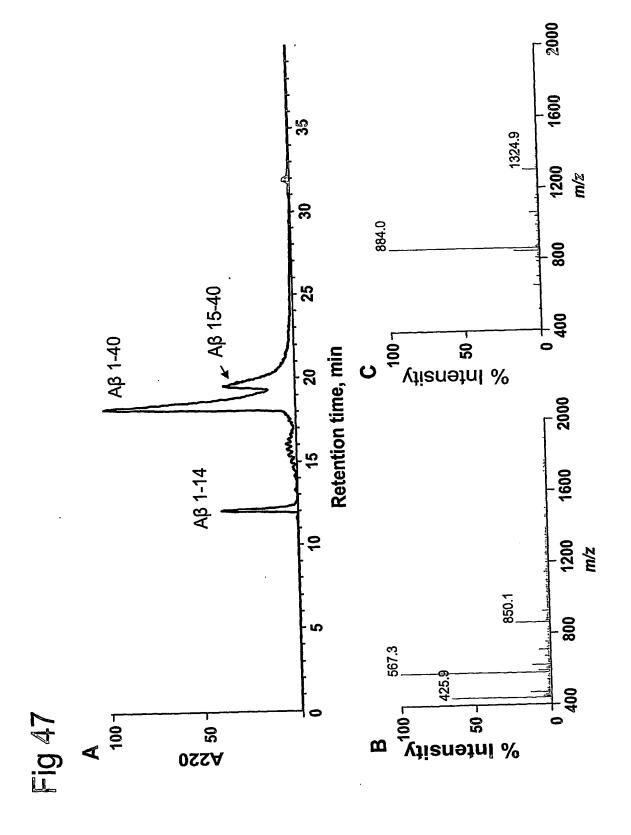








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